

0400

#3

OIPPE

P.5

RAW SEQUENCE LISTING

DATE: 12/14/2001

PATENT APPLICATION: US/10/005,344

TIME: 10:09:38

Input Set : A:\ISPH622_Sequence Listing.txt

Output Set: N:\CRF3\12142001\I005344.raw

ENTERED

```

4 <110> APPLICANT: Loren J. Miraglia
5   Pamela Nero
6   Mark J. Graham
7   Brett P. Monia
8   Erich Koller
9   MingYi Chiang
10  Mano Manoharan
12 <120> TITLE OF INVENTION: Antisense Modulation of mdm2 expression.
14 <130> FILE REFERENCE: ISPH-0622
C--> 16 <140> CURRENT APPLICATION NUMBER: US/10/005,344
C--> 17 <141> CURRENT FILING DATE: 2001-12-04
19 <150> PRIOR APPLICATION NUMBER: US 09/048,810
20 <151> PRIOR FILING DATE: 1998-03-26
22 <150> PRIOR APPLICATION NUMBER: US 09/280,805
23 <151> PRIOR FILING DATE: 1999-03-26
25 <160> NUMBER OF SEQ ID NOS: 379
27 <170> SOFTWARE: FastSEQ for Windows Version 4.0
29 <210> SEQ ID NO: 1
30 <211> LENGTH: 2372
31 <212> TYPE: DNA
32 <213> ORGANISM: Homo sapiens
34 <220> FEATURE:
35 <221> NAME/KEY: CDS
36 <222> LOCATION: (312)...(1787)
38 <400> SEQUENCE: 1
39 gcaccgcgcg agcttggctg cttctggggc ctgtgtggcc ctgtgtgtcg gaaagatgga 60
40 gcaagaagcc gagcccgagg ggcggccgcg acccctctga ccgagatcct gctgctttcg 120
41 cagccaggag caccgtccct ccccgatta gtgcgtacga gcgccagtg ccctggccccg 180
42 gagagtggaa tgatcccga ggcccagggc gtcgtgcttc cgagtagtc agtccccgtg 240
43 aaggaaactg gggagtcttg agggaccccc gactccaagc gcgaaaacc cggatggtga 300
44 ggagcaggca a atg tgc aat acc aac atg tct gta cct act gat ggt gct 350
45           Met Cys Asn Thr Asn Met Ser Val Pro Thr Asp Gly Ala
46           1           5           10
48 gta acc acc tca cag att cca gct tcg gaa caa gag acc ctg gtt aga 398
49 Val Thr Thr Ser Gln Ile Pro Ala Ser Glu Gln Glu Thr Leu Val Arg
50           15           20           25
52 cca aag cca ttg ctt ttg aag tta tta aag tct gtt ggt gca caa aaa 446
53 Pro Lys Pro Leu Leu Leu Lys Leu Leu Lys Ser Val Gly Ala Gln Lys
54 30           35           40           45
56 gac act tat act atg aaa gag gtt ctt ttt tat ctt ggc cag tat att 494
57 Asp Thr Tyr Thr Met Lys Glu Val Leu Phe Tyr Leu Gly Gln Tyr Ile
58           50           55           60
60 atg act aaa cga tta tat gat gag aag caa caa cat att gta tat tgt 542
61 Met Thr Lys Arg Leu Tyr Asp Glu Lys Gln Gln His Ile Val Tyr Cys
62           65           70           75
64 tca aat gat ctt cta gga gat ttg ttt ggc gtg cca agc ttc tct gtg 590
65 Ser Asn Asp Leu Leu Gly Asp Leu Phe Gly Val Pro Ser Phe Ser Val

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66	80	85	90	
68	aaa gag cac agg aaa ata tat acc atg atc tac agg aac ttg gta gta	638		
69	Lys Glu His Arg Lys Ile Tyr Thr Met Ile Tyr Arg Asn Leu Val Val			
70	95 100 105			
72	gtc aat cag cag gaa tca tgc gac tca ggt aca tct gtg agt gag aac	686		
73	Val Asn Gln Gln Glu Ser Ser Asp Ser Gly Thr Ser Val Ser Glu Asn			
74	110 115 120 125			
76	agg tgt cac ctt gaa ggt ggg agt gat caa aag gac ctt gta caa gag	734		
77	Arg Cys His Leu Glu Gly Gly Ser Asp Gln Lys Asp Leu Val Gln Glu			
78	130 135 140			
80	ctt cag gaa gag aaa cct tca tct tca cat ttg gtt tct aga cca tct	782		
81	Leu Gln Glu Glu Lys Pro Ser Ser Ser His Leu Val Ser Arg Pro Ser			
82	145 150 155			
84	acc tca tct aga agg aga gca att agt gag aca gaa gaa aat tca gat	830		
85	Thr Ser Ser Arg Arg Arg Ala Ile Ser Glu Thr Glu Glu Asn Ser Asp			
86	160 165 170			
88	gaa tta tct ggt gaa cga caa aga aaa cgc cac aaa tct gat agt att	878		
89	Glu Leu Ser Gly Glu Arg Gln Arg Lys Arg His Lys Ser Asp Ser Ile			
90	175 180 185			
92	tcc ctt tcc ttt gat gaa agc ctg gct ctg tgt gta ata agg gag ata	926		
93	Ser Leu Ser Phe Asp Glu Ser Leu Ala Leu Cys Val Ile Arg Glu Ile			
94	190 195 200 205			
96	tgt tgt gaa aga agc agt agc agt gaa tct aca ggg acg cca tgc aat	974		
97	Cys Cys Glu Arg Ser Ser Ser Ser Glu Ser Thr Gly Thr Pro Ser Asn			
98	210 215 220			
100	ccg gat ctt gat gct ggt gta agt gaa cat tca ggt gat tgg ttg gat	1022		
101	Pro Asp Leu Asp Ala Gly Val Ser Glu His Ser Gly Asp Trp Leu Asp			
102	225 230 235			
104	cag gat tca gtt tca gat cag ttt agt gta gaa ttt gaa gtt gaa tct	1070		
105	Gln Asp Ser Val Ser Asp Gln Phe Ser Val Glu Phe Glu Val Glu Ser			
106	240 245 250			
108	ctc gac tca gaa gat tat agc ctt agt gaa gaa gga caa gaa ctc tca	1118		
109	Leu Asp Ser Glu Asp Tyr Ser Leu Ser Glu Glu Gly Gln Glu Leu Ser			
110	255 260 265			
112	gat gaa gat gat gag gta tat caa gtt act gtg tat cag gca ggg gag	1166		
113	Asp Glu Asp Asp Glu Val Tyr Gln Val Thr Val Tyr Gln Ala Gly Glu			
114	270 275 280 285			
116	agt gat aca gat tca ttt gaa gaa gat cct gaa att tcc tta gct gac	1214		
117	Ser Asp Thr Asp Ser Phe Glu Glu Asp Pro Glu Ile Ser Leu Ala Asp			
118	290 295 300			
120	tat tgg aaa tgc act tca tgc aat gaa atg aat ccc ccc ctt cca tca	1262		
121	Tyr Trp Lys Cys Thr Ser Cys Asn Glu Met Asn Pro Pro Leu Pro Ser			
122	305 310 315			
124	cat tgc aac aga tgt tgg gcc ctt cgt gag aat tgg ctt cct gaa gat	1310		
125	His Cys Asn Arg Cys Trp Ala Leu Arg Glu Asn Trp Leu Pro Glu Asp			
126	320 325 330			
128	aaa ggg aaa gat aaa ggg gaa atc tct gag aaa gcc aaa ctg gaa aac	1358		
129	Lys Gly Lys Asp Lys Gly Glu Ile Ser Glu Lys Ala Lys Leu Glu Asn			
130	335 340 345			

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```

132 tca aca caa gct gaa gag ggc ttt gat gtt cct gat tgt aaa aaa act 1406
133 Ser Thr Gln Ala Glu Glu Gly Phe Asp Val Pro Asp Cys Lys Lys Thr
134 350 355 360 365
136 ata gtg aat gat tcc aga gag tca tgt gtt gag gaa aat gat gat aaa 1454
137 Ile Val Asn Asp Ser Arg Glu Ser Cys Val Glu Glu Asn Asp Asp Lys
138 370 375 380
140 att aca caa gct tca caa tca caa gaa agt gaa gac tat tct cag cca 1502
141 Ile Thr Gln Ala Ser Gln Ser Gln Glu Ser Glu Asp Tyr Ser Gln Pro
142 385 390 395
144 tca act tct agt agc att att tat agc agc caa gaa gat gtg aaa gag 1550
145 Ser Thr Ser Ser Ser Ile Ile Tyr Ser Ser Gln Glu Asp Val Lys Glu
146 400 405 410
148 ttt gaa agg gaa gaa acc caa gac aaa gaa gag agt gtg gaa tct agt 1598
149 Phe Glu Arg Glu Glu Thr Gln Asp Lys Glu Glu Ser Val Glu Ser Ser
150 415 420 425
152 ttg ccc ctt aat gcc att gaa cct tgt gtg att tgt caa ggt cga cct 1646
153 Leu Pro Leu Asn Ala Ile Glu Pro Cys Val Ile Cys Gln Gly Arg Pro
154 430 435 440 445
156 aaa aat ggt tgc att gtc cat ggc aaa aca gga cat ctt atg gcc tgc 1694
157 Lys Asn Gly Cys Ile Val His Gly Lys Thr Gly His Leu Met Ala Cys
158 450 455 460
160 ttt aca tgt gca aag aag cta aag aaa agg aat aag ccc tgc cca gta 1742
161 Phe Thr Cys Ala Lys Lys Leu Lys Lys Arg Asn Lys Pro Cys Pro Val
162 465 470 475
164 tgt aga caa cca att caa atg att gtg cta act tat ttc ccc tag 1787
165 Cys Arg Gln Pro Ile Gln Met Ile Val Leu Thr Tyr Phe Pro *
166 480 485 490
168 ttgacctgtc tataagagaa ttatatattt ctaactatat aaccctagga atttagacaa 1847
169 cctgaaattt attcacatat atcaaagtga gaaaatgcct caattcacat agatttcttc 1907
170 cttttagtagt aattgacctt ctttggtagt ggaatagtagt atacttacta taatttgact 1967
171 tgaatatgta gctcatcctt tacaccaact cctaatttta aataatttct actctgtctt 2027
172 aaatgagaag tacttggttt tttttttctt aaatatgtat atgacattta aatgtaactt 2087
173 attatttttt ttgagaccga gtcttgctct gttaccacagg ctggagtgca gtgggtgatc 2147
174 ttggctcact gcaagctctg cctcccccg gttcgaccca ttctcctgcc tcagcctccc 2207
175 aattagcttg gctacagtc atctgccacc acacctggct aattttttgt acttttagta 2267
176 gagacagggg ttcaccgtgt tagccaggat ggtctcgatc tcctgacctc gtgatccgcc 2327
177 cacctcggcc tccaaagtg ctgggattac aggcattgagc caccg 2372
179 <210> SEQ ID NO: 2
180 <211> LENGTH: 500
181 <212> TYPE: DNA
182 <213> ORGANISM: Homo sapiens
184 <220> FEATURE:
185 <221> NAME/KEY: misc_signal
186 <222> LOCATION: (138)...(157)
187 <223> OTHER INFORMATION: p53 response element RE1
189 <221> NAME/KEY: misc_signal
190 <222> LOCATION: (176)...(195)
191 <223> OTHER INFORMATION: p53 response element RE2
193 <221> NAME/KEY: exon

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194 <222> LOCATION: (231)...(301)
195 <223> OTHER INFORMATION: Exon 2
197 <221> NAME/KEY: intron
198 <222> LOCATION: (302)...(422)
199 <223> OTHER INFORMATION: Intron 2
201 <400> SEQUENCE: 2
202 ggctgcgggc ccctgcggcg cgggaggtcc ggatgatcgc aggtgcctgt cgggtcacta 60
203 gtgtgaacgc tgcgcgtagt ctgggcggga ttgggccggg tcagtgggca ggttgactca 120
204 gcttttcctc ttgagctggt caagttcaga cacgttcoga aactgcagta aaaggagtta 180
205 agtcctgact tgtctccagc tggggctatt taaacatgc attttcccag ctgtgttcag 240
206 tggcgattgg agggtagacc tgtgggcacg gacgcacgcc actttttctc tgctgatcca 300
207 ggtaagcacc gacttgcttg tagctttagt ttttaactgtt gtttatgttc tttatatatg 360
208 atgtattttc cacagatggt tcatgatttc cagttttcat cgtgtctttt ttttccttgt 420
209 aggcaaatgt gcaataccaa catgtctgta cctactgatg gggctgtaac caccacacag 480
210 attccagctt cggaacaaga                               500
212 <210> SEQ ID NO: 3
213 <211> LENGTH: 20
214 <212> TYPE: DNA
215 <213> ORGANISM: Artificial Sequence
217 <220> FEATURE:
218 <223> OTHER INFORMATION: Antisense Oligonucleotide
220 <400> SEQUENCE: 3
221 cagccaagct cgcgcgggtgc                               20
223 <210> SEQ ID NO: 4
224 <211> LENGTH: 20
225 <212> TYPE: DNA
226 <213> ORGANISM: Artificial Sequence
228 <220> FEATURE:
229 <223> OTHER INFORMATION: Antisense Oligonucleotide
231 <400> SEQUENCE: 4
232 tctttccgac acacagggcc                               20
234 <210> SEQ ID NO: 5
235 <211> LENGTH: 20
236 <212> TYPE: DNA
237 <213> ORGANISM: Artificial Sequence
239 <220> FEATURE:
240 <223> OTHER INFORMATION: Antisense Oligonucleotide
242 <400> SEQUENCE: 5
243 cagcaggatc tcggtcagag                               20
245 <210> SEQ ID NO: 6
246 <211> LENGTH: 20
247 <212> TYPE: DNA
248 <213> ORGANISM: Artificial Sequence
250 <220> FEATURE:
251 <223> OTHER INFORMATION: Antisense Oligonucleotide
253 <400> SEQUENCE: 6
254 gggcgctcgt acgcactaat                               20
256 <210> SEQ ID NO: 7
257 <211> LENGTH: 20

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Input Set : A:\ISPH622_Sequence Listing.txt

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```

258 <212> TYPE: DNA
259 <213> ORGANISM: Artificial Sequence
261 <220> FEATURE:
262 <223> OTHER INFORMATION: Antisense Oligonucleotide
264 <400> SEQUENCE: 7
265 tcggggatca ttccactctc 20
267 <210> SEQ ID NO: 8
268 <211> LENGTH: 20
269 <212> TYPE: DNA
270 <213> ORGANISM: Artificial Sequence
272 <220> FEATURE:
273 <223> OTHER INFORMATION: Antisense Oligonucleotide
275 <400> SEQUENCE: 8
276 cgggggttttc gcgcttgag 20
278 <210> SEQ ID NO: 9
279 <211> LENGTH: 20
280 <212> TYPE: DNA
281 <213> ORGANISM: Artificial Sequence
283 <220> FEATURE:
284 <223> OTHER INFORMATION: Antisense Oligonucleotide
286 <400> SEQUENCE: 9
287 catttgcctg ctccctcacca 20
289 <210> SEQ ID NO: 10
290 <211> LENGTH: 20
291 <212> TYPE: DNA
292 <213> ORGANISM: Artificial Sequence
294 <220> FEATURE:
295 <223> OTHER INFORMATION: Antisense Oligonucleotide
297 <400> SEQUENCE: 10
298 gtattgcaca ttgacctgct 20
300 <210> SEQ ID NO: 11
301 <211> LENGTH: 20
302 <212> TYPE: DNA
303 <213> ORGANISM: Artificial Sequence
305 <220> FEATURE:
306 <223> OTHER INFORMATION: Antisense Oligonucleotide
308 <400> SEQUENCE: 11
309 agcaccatca gtaggtacag 20
311 <210> SEQ ID NO: 12
312 <211> LENGTH: 20
313 <212> TYPE: DNA
314 <213> ORGANISM: Artificial Sequence
316 <220> FEATURE:
317 <223> OTHER INFORMATION: Antisense Oligonucleotide
319 <400> SEQUENCE: 12
320 ctaccaagtt cctgtagatc 20
322 <210> SEQ ID NO: 13
323 <211> LENGTH: 20
324 <212> TYPE: DNA

```

→ Use of n and / or Xaa has been detected in the Sequence Listing. Review the Sequence Listing to ensure a corresponding explanation is present in the <220> to <223> fields of each sequence using n or Xaa.

VERIFICATION SUMMARY

DATE: 12/14/2001

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Input Set : A:\ISPH622_Sequence Listing.txt

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L:16 M:270 C: Current Application Number differs, Replaced Application Number
L:17 M:271 C: Current Filing Date differs, Replaced Current Filing Date
L:4424 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:370
L:4425 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:370
L:4536 M:334 W: (2) Invalid Amino Acid in Coding Region, NUMBER OF INVALID KEYS:3